Amendments to the Claims:

This listing of claims will replace all prior versions and listings of claims in the application. Please amend claims 80-88.

Listing of Claims:

- 1. 79. (Canceled)
- 80. (Currently Amended) A-ligand-activated uni-molecular detector An polypeptide comprising consisting essentially of:

a circularly permutated marker protein comprising a first interactor domain covalently bonded to the a circularly permutated marker protein through an N-terminal breakpoint of the circularly permutated marker protein and a second interactor domain covalently bonded to the circularly permutated marker protein through a C-terminal breakpoint of the circularly permutated marker protein, wherein said circularly permutated marker protein is functionally reconstituted only upon binding of said first interactor domain and said second interactor domain to a single ligand.

- 81. (Currently Amended) The ligand-activated uni-molecular detector polypeptide of claim 80, wherein said circularly permutated marker protein is a circularly permutated enzyme.
- 82. (Currently Amended) The **ligand-activated uni-molecular detector polypeptide** of claim 81, wherein said circularly permutated enzyme is a β-lactamase protein.
- 83. (Currently Amended) The ligand-activated uni-molecular detector **polypeptide** of claim 82, wherein said circularly permutated enzyme is a TEM-1 β-lactamase protein.

84. (Currently Amended) The ligand-activated uni-molecular detector polypeptide of claim 80, wherein said N-terminal break point and said C-terminal break point are within a solvent exposed loop between elements of secondary structure within the enzyme.

85. (Currently Amended) The ligand-activated uni-molecular detector polypeptide of claim 80, wherein said eircularly permutated protein consists essentially of a circularly permutated marker protein is a β-lactamase protein that has been circularly permutated, with the said β-lactamase protein consisting of amino acids 26 to 288 of the following sequence prior to circular permutation following numbering convention:

His Pro Glu Thr Leu Val Lys Val Lys Asp Ala Glu Asp Gln Leu Gly Ala Arg Val Gly Tyr Ile Glu Leu Asp Leu Asn Ser Gly Lys Ile Leu Glu Ser Phe Arg Pro Glu Glu Arg Phe Pro Met Met Ser Thr Phe Lys Val Leu Cys Gly Ala Val Leu Ser Arg Ile Asp Ala Gly Gln Glu Gln Leu Gly Arg Arg Ile His Tyr Ser Gln Asn Asp Leu Val Glu Tyr Ser Pro Val Thr Glu Lys His Leu Thr Asp Gly Met Thr Val Arg Glu Leu Cys Ser Ala Ala Ile Thr Met Ser Asp Asn Thr Ala Ala Asn Leu Leu Leu Thr Thr Ile Gly Gly Pro Lys Glu Leu Thr Ala Phe Leu His Asn Met Gly Asp His Val Thr Arg Leu Asp Arg Trp Glu Pro Glu Leu Asn Glu Ala Ile Pro Asn Asp Glu Arg Asp Thr Thr Met Pro Val Ala

Met Ala Thr Thr Leu Arg Lys Leu Leu Thr Gly Glu Leu Leu Thr Leu Ala Ser Arg Gln Gln Leu Ile Asp Trp Met Glu Ala Asp Lys Val Ala Gly Pro Leu Leu Arg Ser Ala Leu Pro Ala Gly Trp Phe Ile Ala Asp Lys Ser Gly Ala Gly Glu Arg Gly Ser Arg Gly Ile Ile Ala Ala Leu Gly Pro Asp Gly Lys Pro Ser Arg Ile Val Val Ile Tyr Thr Thr Gly Ser Gln Ala Thr Met Asp Glu Arg Asn Arg Gln Ile Ala Glu Ile Gly Ala Ser Leu Ile Lys His Trp

(SEQ ID NO: 2);

wherein said N-terminal breakpoint and said C-terminal breakpoint are within 10 amino acids of an amide bond junction between two amino acids selected from the group consisting of asparagine 52 and serine 53, leucine 91 and glycine 92, glutamine 99 and asparagine 100, proline 174 and asparagine 175, glutamic acid 197 and leucine 198, lysine 215 and valine 216, alanine 227 and glycine 228, and glycine 253 and lysine 254.

- 86. (Currently Amended) The ligand-activated-uni-molecular detector polypeptide of claim 85, wherein said two amino acids are selected from the group consisting of proline 174 and asparagine 175, glutamic acid 197 and leucine 198, lysine 215 and valine 216, alanine 227 and glycine 228, and glycine 253 and lysine 254.
- 87. (Currently Amended) The ligand-activated-uni-molecular detector polypeptide of claim 85, wherein said two amino acids are glutamic acid 197 and leucine 198.

88. (Currently Amended) The ligand-activated uni-molecular detector **polypeptide** of claim 80, wherein said ligand is a protein ligand.